

Virus Name: Prospect Hill		Abbreviation: PHV
Status Probably not Arbovirus	Select Agent No	SALS Level
SALS Basis		
Other Information		
Antigenic Group Hantaan		

SECTION I - Full Virus Name and Prototype Number

Prototype Strain Number / Designation MP-40	Accession Number	Original Date Submitted 5/31/1985
Family Bunyaviridae	Genus Hantavirus	
Information From P. W. Lee and R. Yanagihara	Address Laboratory of Central Nervous System Studies, National Institute of Neurological and Communicative Disorders and Stroke, Bidg.	
Information Footnote Reviewed by editor		

Section II - Original Source

Isolated By (name) Pyung-Woo Lee, et al (1)	Isolated at Institute Frederick, Maryland	
Host Genus Microtus pennsylvanicus (meadow vole)	Species	Host Age/Stage Adult
Sex Not Answered		
<u>Isolated From</u>	<u>Isolation Details</u>	
Organs/Tissues	Lung	
Signs and Symptoms of Illness None (subclinical infection)	Arthropod	
Time Held Alive before Inoculation		
Collection Method Live caught (Sherman trap)	Collection Date 9/4/1982	
Place Collected (Minimum of City, State, Country) Prospect Hill, Frederick, Maryland, USA		
Latitude 39° 25' N	Longitude 77° 21' W	
Macrohabitat Meadow	Microhabitat	Method of Storage until Inoculated Kept alive until processed
Footnotes		

Section III - Method of Isolation

Inoculation Date
9/27/1982

Animal (Details will be in Section 6)
Microtus, E6 (Vero)

Route Inoculated Intramuscular	Reisolation Yes
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Other Reasons

Homologous Antibody Formation by Source Animal
Yes

Test(s) Used
Indirect IFA

Footnotes

Section IV - Virus Properties

Physicochemical
RNA, Single Strand

Pieces (number of genome segments)	Infectivity	Sedimentation Coefficients(s) (S)
Percentage wt, of Virion Protein	Lipid	Carbohydrate
Virion Polypeptides: Number	Details	
Non-virion Polypeptides: Number	Details	
Virion Density	Sedimentation Coefficients(s) (S)	
Nucleocapsid Density	Sedimentation Coefficients(s) (S)	

Stability of Infectivity (effects)

pH (infective range)
Stable at pH 7.4; inactivated at pH 3.0

Lipid Solvent (ether - % used to test) 20%	After Treatment Titer <1.0 dex	Control Titer 4.5 dex
Lipid Solvent (chloroform) 5%	After Treatment Titer <1.0 dex	Control Titer 4.5 dex
Lipid Solvent (deoxycholate) 0.2%	After Treatment Titer <1.0 dex	Control Titer 4.5 dex

Other (formalin, radiation)
Inactivated by ultraviolet irradiation and heat (56dC; 30 min.)

Virion Morphology

Shape	Dimensions	
Mean 102 nmnm	Range 80-120 nmnm	
Measurement Method Ultrafiltration	Surface Projections/Envelope	Nucleocapsid Dimensions, Symmetry

Morphogenesis

Site of Constituent Formation in Cell	Site of Virion Assembly	Site of Virion Accumulation
Inclusion Bodies None demonstrable	Other	

Hemagglutination

Hemagglutination yes, not t	Antigen Source PEG conc. of E6 cell culture fluid	Erythrocytes (species used)
pH Range 5.8-7.2	pH Optimum 5.8	
Temperature Range	Temperature Optimum Only RT tried	

Remarks

Serologic Methods Recommended
IFA, ELISA, PRNT, immune adherence HA

Footnotes

Section V - Antigenic Relationship and Lack of Relationship to Other Viruses

Serological relationships between Prospect Hill virus, Hantaan (HTN) virus and nephropathia epidemica (NE) virus were investigated using the indirect immunofluorescent antibody (IFA) technique. Sera were tested simultaneously against all three antigens, prepared as virus-infected E6 cells. The resulting antibody profiles were sufficiently distinct to permit discrimination between these viruses by IFA. In addition:

1. Sera from four American mammalogists and sera from meadow voles trapped in the United States reacted to highest titer against Prospect Hill virus and 2- to 4-fold and 4- to 8-fold, respectively, less to NE and HTN viruses.
2. Sera from patients with classical, severe hemorrhagic fever with renal syndrome (HFRS) in Korea, Far Eastern USSR, and the People's Republic of China reacted to highest titer to HTN virus and 64- to 128-fold and 128- to 256-fold, respectively, lower to Prospect Hill virus and NE virus.
3. Sera from the majority of patients with HFRS in Scandinavia, European USSR (Tula and Bashkiria), Hungary, and Yugoslavia reacted to highest titer to NE virus and 2-fold and 4-fold, respectively, lower to Prospect Hill virus and HTN virus.
4. Six monoclonal antibodies produced to HTN virus were used to examine antigenic relationships by IFA between HTN, Prospect Hill and NE viruses. All six monoclonal antibodies reacted with HTN virus, but none reacted with Prospect Hill or NE viruses [2].

Section VI - Biologic Characteristics

Virus Source (all VERTEBRATE isolates)
Lung (LV), blood (LV)

Lab Methods of Virus Recovery (ALL ISOLATIONS)
Intramuscular inoculation of seronegative, laboratory-bred suckling or weanling *Microtus pennsylvanicus*; E6 clone of Vero cells

Cell system (a)	Virus passage history (b)	Evidence of Infection						
		CPE			PLAQUES			Growth Without CPE +/- (g)
		Day (c)	Extent (d)	Titer TCD50/ml (e)	Day (c)	Size (f)	Titer PFU/ml (e)	
E6 (CL)	None	10	2+	3.0*				
E6 (CL)	E6/P5	2-29	1+-4+	2.0-5.0				
E6 (CL)	E6/P11				10	5mm	7.2*	

* Expressed in dex

Vertebrate (species and organ) and arthropod	No. isolations/No. tested	No. with antibody/No. tested Test used	Country and region
Microtus pennsylvanicus	2/14	53 [*] /230 IFA	Pennsylvania; Alaska, USA
Microtus californicus		46/185 IFA	California, USA
Microtus montanus		0/15 IFA	
Microtus oeconomus		0/14 IFA	Alaska, USA
Microtus breweri		0/12 IFA	Massachusetts, USA
Microtus oregoni		0/2 IFA	California, USA
Peromyscus maniculatus		5/41 IFA	California; Minnesota, USA
Peromyscus truei		1/18 IFA	California, USA
Peromyscus leucopus		0/220 IFA	Maryland; Virginia Pennsylvania, USA
Peromyscus eremicus		0/17 IFA	California, USA
Peromyscus crinatus		0/17 IFA	
Peromyscus boylii		0/3 IFA	
Clethrionomys gapperi		0/11 IFA	Pennsylvania; Minnesota, USA
Clethrionomys rutilus		0/9 IFA	
Neotoma micropus		0/25 IFA	Texas, USA
Humans (mammalogists)		4/203 IFA,NT	U.S.A. (4)

* Serum samples having immunofluorescent antibody titers of >16 to Prospect Hill virus were considered presumptive evidence of previous infection.

Experimental host and age	Passage history and strain	Inoculation Route-Dose	Evidence of infection	AST (days)	Titer log ₁₀ /ml
Balb/c mice (nb)	MP-40 strain, E6/P5	ic 0.01	Develop antibody (IFA)		
Mice (nb)		ip			
Mice (nb)		sc			
Mice (wn)		ic			
Mice (wn)		ip			
Mice (wn)		im 0.1	Develop antibody (IFA)		
Nude mice (NIH:S)		im 0.1	No antibody or antigen		
Fischer rat (nb)		ic 0.01	Develop antibody (IFA)		
Fischer rat (wn)		im 0.3	Develop antibody (IFA)		
Syrian hamster (wn)		im 0.3	Develop antibody (IFA)		
Mongolian gerbil (wn)		im 0.3	Develop antibody (IFA)		
Meadow vole (nb)		ic 0.01	Specific antibody and antigen		
Meadow vole (wn)		im 0.3	in lung tissue 14 days pi		
Cynomolgus monkey		iv 10 ml	Viremia day 7, specific antibody; mild proteinuria		
Cynomologus monkeys		iv	mild, transient proteinuria and azotemia (5)		
Chimpanzee	iv	Mild, transient proteinuria and azotemia (5)			

Section IX - Experimental Arthropod Infection and Transmission

Arthropod species & virus source(a)	Method of Infection log10/ml (b)		Incubation period (c)		Transmission by bite (d)		Assay of arthropod, log10/ml (e)		
	Feeding	Injected	Days	°C	Host	Ratio	Whole	Organ	System

Section X - Histopathology

Character of lesions (specify host)

No significant lesion found in naturally and experimentally infected meadow voles.

Inclusion Bodies

Intranuclear

Organs/Tissues Affected

Category of tropism

Section XI - Human Disease

In Nature

Residual

Death

Subclinical

Overt Disease

Clinical Manifestations

Number of Cases

Category (i.e. febrile illness, etc.)

Section XII - Geographic Distribution

Known (Virus detected)

Maryland, USA

Suspected (Antibody only detected)

Alaska; California; Virginia, USA.

Section XIII - References

1. Lee, P.-W., et al. 1982. *Lancet* ii(8312):1405-1407.
2. Franko, M.C., et al. 1983. *Proc. Nat. Acad. Sci.* 80:4149-4153.
3. Tsai, T.F. Personal communication. 1984.
4. Yanigihara, R. et al. 1984. *New England J. Med.* 310:1325-1326".
5. Yanagihara, R. et al. 1988. *Arch. virol.* 101:125-130.

Remarks